10/049,511 Page 3

L1 HAS NO ANSWERS L1 STR



Structure attributes must be viewed using STN Express query preparation.

=> s l1 SAMPLE SEARCH INITIATED 15:23:30 FILE 'REGISTRY' SAMPLE SCREEN SEARCH COMPLETED - 14 TO ITERATE

100.0% PROCESSED 14 ITERATIONS 3 ANSWERS

SEARCH TIME: 00.00.01

FULL FILE PROJECTIONS: ONLINE **COMPLETE**

BATCH **COMPLETE**

PROJECTED ITERATIONS: 56 TO 504

PROJECTED ANSWERS: 3 TO 163

L2 3 SEA SSS SAM L1

=> s 11 sss full FULL SEARCH INITIATED 15:23:40 FILE 'REGISTRY'

FULL SCREEN SEARCH COMPLETED - 300 TO ITERATE

100.0% PROCESSED 300 ITERATIONS 20 ANSWERS

SEARCH TIME: 00.00.01

L3 20 SEA SSS FUL L1

=> file caplus

COST IN U.S. DOLLARS SINCE FILE TOTAL

ENTRY SESSION

FULL ESTIMATED COST 148.36

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FILE COVERS 1907 - 9 May 2003 VOL 138 ISS 20 FILE LAST UPDATED: 8 May 2003 (20030508/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s 132 L3 L4

=> d ibib abs hitstr tot

ANSWER 1 OF 2 CAPLUS COPYRIGHT 2003 ACS ACCESSION NUMBER: 2001:137218 CAPLUS

DOCUMENT NUMBER:

134:193607

TITLE:

Preparation of phenanthrolin-7-one derivatives and

their therapeutic uses as antitumoral medicines Delfourne, Evelyne; Darro, Francis; Bastide, Jean;

Kiss, Robert; Frydman, Armand

PATENT ASSIGNEE(S):

Laboratoire L. Lafon, Fr. PCT Int. Appl., 54 pp.

SOURCE:

INVENTOR(S):

CODEN: PIXXD2

DOCUMENT TYPE:

Patent French

LANGUAGE: FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

													_					
PATENT NO. KI					ND	DATE		APPLICATION NO.						DATE				
WO									WO 2000-FR2313					20000811				
WO	2001012632			A3		20010719												
	W:	ΑE,	AG,	AL,	AM,	ΑT,	ΑU,	AZ,	BA,	BB,	BG,	BR,	BY,	BZ,	CA,	CH,	CN,	
		CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	ЕĒ,	ES,	FI,	GB,	GD,	GE,	GH,	GM,	HR,	
		HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	ΚP,	KR,	ΚZ,	LC,	LK,	LR,	LS,	LT,	
														PL,				
		SD,	SE,	SG,	SI,	SK,	SL,	ТJ,	TM,	TR,	TT,	TZ,	UA,	ŪĠ,	US,	UΖ,	VN,	
						AZ,												
	RW:													ΑT,	BE,	CH,	CY,	
		DE,	DK.	ES,	FI,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	BF,	ВJ,	
		CF,	CG,	CI,	CM,	GA,	GN,	GW,	ML,	MR,	NE,	SN,	TD,	TG				
FR	2797446			A1 20010216				FR 1999-10493					19990813					
FR	2797446				B1 20011102													
BR	2000013239			A	A 2		20020423		BR 2000-13239 20000811									
									EP 2000-958679 20000811									
														NL,		MC,	PT,	
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NO	2002	0006	69	, A	,	2002	0415	•	N	0 20	02-6	69		2002	0211			
PRIORITY APPLN. INFO.:									FR 1999-10493				Α	19990813				
INIONII							WO 2000-FR2313											
OTHER SOURCE(S): CASREACT 134:193607; MARPAT 134:193607																		

5/9/2003

GI

The invention concerns a pharmaceutical compn. comprising an efficient AB amt. of a compd. selected among the compds. I [R1, R2, R3, R4, R5 = H, halogen, C1-6-alkyl, OH, CHO, OR8, CO2H, CN, CO2R8, CONHR8, CONR8R9, NH2, NHR8, N(R8)2, NHCH2CH2NMe2, NHCH2CH2Cl, NHCOR8, morpholino, NO2, SO3H, CH2N(CO2R8)CH2CO2R9, CH2N(CO2R8)CH2Ar; R6 = H, halogen, C1-6-alkyl, (CH2) nR10,; R7 = H, C1-6-alkyl, Ph-C1-4-alkyl, NR15R16; R8, R9 = C1-6-alkyl, Ph-C1-4-alkyl; R10 = halogen, OH, C1-6-alkoxy, OC(:0)-C1-6-alkyl, CN, CO2Et, COR11; R11 = Ph-C1-4-alkyl, NR12R13; R12 , R13 = H, C1-6-alkyl, Ph-C1-4-alkyl, (CH2)nR14; R14 = halogen, C1-6-alkoxy, NMe2; R15, R16 = H, C1-6-alkyl, Ph-C1-4-alkyl, (CH2) nR17; R17 = H, halogen, OH, C1-6-alkoxy; Ar = C6-14-aryl; n = 1 - 6 and II or their pharmaceutically acceptable salts. Thus, I [R1 = R2 = R3 = R4 = R5 = R6 = R7 = H (CRL8293)] and II [R1 = R2 = R3 = R4 = R5 = R6 = R7 = H (CRL8294)] were prepd. from quinoline-5,8-dione via Diels-Alder with crotonaldehyde dimethylhydrazone followed by cyclocondensation of the resulting quinone III with Me2NCMe(OEt)2. I (R1 = R2 = R3 = R4 = R5 = R6 = R7 = H) and II (R1 = R2 = R3 = R4 = R5 = R6 = R7 = H) have interesting cytotoxic properties [DMT = 10 mg/Kg (DMT = max. tolerable dose); -33% and -36%, resp. tumor surface diminution {murin mammary carcinoma (MXT-HI)}; -45% and -64% , resp. tumor surface diminution [{murin mammary adenocarcinoma (MXT-HS)]; and, for II, T/C = 136% (lymphoma L1210)] leading to a therapeutic use as antitumoral medicines.

THE TAPEUTIC USE AS ARCTIUMOTAL MEDICINES.

1T 266306-76-7P, CRL 8294 327184-13-4P, CRL 8364

327184-14-5P 327184-15-6P, CRL 8401 327184-16-7P

, CRL 8440 327184-17-8P, CRL 8479 327184-18-9P

327184-19-0P, CRL 8485 327184-20-3P 327184-21-4P

3-(Acetoxymethyl)-9-methoxy-7H-pyrido[4,3,2-de][1,10]phenanthrolin-7-one

327184-22-5P, CRL 8830 327184-33-8P, CRL 8367

327184-35-0P, CRL 8388 327184-37-2P, CRL 8441 327184-39-4P, CRL 8482 327184-41-8P, CRL 8483

327184-39-4P, CRL 8482 327184-41-8P, CRL 8483 327184-43-0P, CRL 8486 327184-45-2P, CRL 8487

327184-43-0P, CRL 8486 327184-45-2P, CRL 8487 327184-47-4P, CRL 8480 327184-49-6P, CRL 8481

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(prepn. of phenanthrolin-7-one derivs. and their therapeutic uses as antitumoral medicines)

RN 266306-76-7 CAPLUS

CN 7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one (9CI) (CA INDEX NAME)

RN 327184-13-4 CAPLUS

CN 7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 11-methoxy- (9CI) (CA INDEX NAME)

RN 327184-14-5 CAPLUS

CN 7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 11-chloro- (9CI) (CA INDEX NAME)

RN 327184-15-6 CAPLUS

CN 7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 4-methoxy- (9CI) (CA INDEX NAME)

RN 327184-16-7 CAPLUS

CN 7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 4,11-dimethoxy- (9CI) (CA INDEX NAME)

RN 327184-17-8 CAPLUS

CN 7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 4,9-dimethoxy- (9CI) (CA INDEX NAME)

RN 327184-18-9 CAPLUS

CN 7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 9-methoxy- (9CI) (CA INDEX NAME)

RN 327184-19-0 CAPLUS

CN 7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 9,11-dimethoxy- (9CI) (CA INDEX NAME)

RN 327184-20-3 CAPLUS

CN 7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 3-[(acetyloxy)methyl]- (9CI)

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(CA INDEX NAME)

RN 327184-21-4 CAPLUS

CN 7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 3-[(acetyloxy)methyl]-9-methoxy- (9CI) (CA INDEX NAME)

RN 327184-22-5 CAPLUS

CN 7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 2-(2-chloroethyl)- (9CI) (CA INDEX NAME)

RN 327184-33-8 CAPLUS

CN 7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 11-(dimethylamino)- (9CI) (CA INDEX NAME)

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327184-35-0 CAPLUS RN

7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 11-hydroxy- (9CI) (CA INDEX CN

327184-37-2 CAPLUS RN

7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 11-(dimethylamino)-4-methoxy-CN (9CI) (CA INDEX NAME)

327184-39-4 CAPLUS RN

7H-Pyrido[4,3,2-de][1,7]phenanthroline-10-carboxylic acid, 7-oxo-, ethyl CN ester (9CI) (CA INDEX NAME)

327184-41-8 CAPLUS RN

7H-Pyrido[4,3,2-de][1,7]phenanthroline-7,9(8H)-dione (9CI) (CA INDEX CN

RN 327184-43-0 CAPLUS

CN 7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 9-chloro-11-(dimethylamino)-(9CI) (CA INDEX NAME)

RN 327184-45-2 CAPLUS

CN 7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 4-hydroxy-, dihydriodide (9CI) (CA INDEX NAME)

●2 HI

RN 327184-47-4 CAPLUS

CN 7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 4-chloro- (9CI) (CA INDEX NAME)

327184-49-6 CAPLUS RN

7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one, 4-(dimethylamino)- (9CI) (CA CN INDEX NAME)

ANSWER 2 OF 2 CAPLUS COPYRIGHT 2003 ACS

ACCESSION NUMBER: DOCUMENT NUMBER:

2000:177139 CAPLUS

132:303121

TITLE:

Mechanism of action studies of cytotoxic marine alkaloids: ascididemin exhibits thiol-dependent

oxidative DNA cleavage

AUTHOR (S):

Matsumoto, Sandra S.; Sidford, Mathew H.; Holden,

Joseph A.; Barrows, Louis R.; Copp, Brent R.

CORPORATE SOURCE:

Departments of Pharmacology and Toxicology, University

of Utah, Salt Lake City, UT, 84112, USA

SOURCE:

Tetrahedron Letters (2000), 41(10), 1667-1670

CODEN: TELEAY; ISSN: 0040-4039

PUBLISHER:

Elsevier Science Ltd.

DOCUMENT TYPE:

Journal

LANGUAGE:

English

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The cytotoxic marine alkaloid ascididemin has been shown to be a thiol-dependent DNA cleaving agent. Previous mechanisms of action studies have concluded that DNA and/or the DNA processing enzyme topoisomerase II were the cellular targets for the alkaloid - this is the first direct evidence that a pyridoacridone alkaloid can cause DNA cleavage under physiol. conditions.

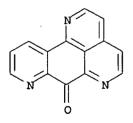
266306-76-7P IT

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(cytotoxic ascididemin exhibits thiol-dependent oxidative DNA cleavage)

266306-76-7 CAPLUS RN

7H-Pyrido[4,3,2-de][1,7]phenanthrolin-7-one (9CI) (CA INDEX NAME) CN



REFERENCE COUNT:

THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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5/9/2003